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ISIC0009-101 (CHEM0005US.P1)

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Attorney Docket Number

| U.S. PATENT DOCUMENTS | | | | | | | |
|------------------------|------|--|------------------------|----------------------------------|---------------------------------------|--|--|
| Examiner Initials * | Cite | Document Number | Publication/Issue Date | Name of Patentee or Applicant of | Pages, Columns, Lines, Where Relevant | | |
| | No.1 | Number - Kind Code ² (if known) | MM-DD-YYYY | Cited Document | Passages or Relevant Figures Appear | | |
| | HG | US-5,998,588 | 12-07-1999 | Hoffman et al. | | | |
| | HI | US-2003/0190635 A1 | 10-09-2003 | McSwiggen et al. | | | |
| | HJ | US-2002/0102267 A1 | 08-01-2002 | Lu et al. | | | |
| | HK | US-2002/0151512 A1 | 10-17-2002 | Peyman et al. | | | |
| | HL | US-2002/0156235 A1 | 10-24-2002 | Manoharan et al. | | | |
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| FOREIGN PATENT DOCUMENTS | | | | | | | |
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| Examiner Initials* | Cite No. ¹ | Foreign Patent Document | Publication | Name of Patentee or Applicant of Cited Document | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear | | |
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| INITO | | DIC | OL OCUPE | Application Number | 10/700,971 | |
| | | | CLOSURE | Filing Date | November 4, 2003 | |
| STATEMENT BY APPLICANT | | | | First Named Inventor | Muthiah Manoharan | |
| | | | | Art Unit | To Be Determined | |
| (Use as many sheets as necessary) | | | necessary) | Examiner Name | To Be Determined | |
| Sheet | 2 | of | 2 | Attorney Docket Number | ISIC0009-101 (CHEM0005US.P1) | |

| NON PATENT LITERATURE DOCUMENTS | | | | | | |
|---------------------------------|--------------------------|---|----|--|--|--|
| Examiner Initials * | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published. | T² | | | |
| | НМ | Astriab-Fisher et al., "Conjugates of antisense oligonucleotides with the TAT and antennapedia cell-penetrting peptides: effects on cellular update, binding to target sequences and biologic actions," <i>Pharmaceutical Research</i> (2002) 19(6):744-754. | | | | |
| | HN | Fire et al., "RNA-triggered gene silencing," TIG (1999) 15(9):358-363. | | | | |
| | НО | Veronese et al., "Bioconjugation in pharmaceutical chemistry," IL Farmaco (1999) 54:497-516. | | | | |
| | НР | International Search Report dated March 24, 2005 for International Application No. PCT/US03/35088. | _ | | | |
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| Examiner Signature | /Sean Mcgarry/ | Date Considered | 06/16/2008 |
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